Attorney's Docket No.: 14879-090002 / D1-A0001YIP-Applicant: Masatake Kudoh et al.

USD1

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## Amendments to the Claims:

This listing of claims replaces all prior versions and listings of claims in the application:

## 1.-14. (Canceled)

- 15. (Currently amended) A method for producing a alcohol an (S)-4-halo-3hydroxybutyric acid ester derivative, the method comprising reacting an (R)-2-octanol dehydrogenase having a molecular weight of about 30,000 Da as determined by SDS-PAGE and about 83,000 Da as determined by gel filtration, or a microorganism producing the (R)-2-octanol dehydrogenase, or a processed product of the microorganism with a 4-haloacetoacetic acid ester derivative-ketone to reduce the 4-haloacetoacetic acid ester derivative-ketone, wherein the (R)-2octanol dehydrogenase is a polypeptide selected from the group from (a) to (e) below:
- (a) a polypeptide encoded by a polynucleotide that hybridizes under stringent conditions with a nucleic acid probe consisting of the complement of the nucleotide sequence of SEQ ID NO: 1;
- (b) a polypeptide comprising an amino acid sequence at least 70% identical to the amino acid sequence of SEQ ID NO:2;
  - (c) a polypeptide comprising the amino acid sequence of SEQ ID NO:2;
- (d) a polypeptide comprising the amino acid sequence of SEQ ID NO:2 with up to 50 conservative amino acid substitutions; and
- (e) a polypeptide comprising the amino acid sequence of SEQ ID NO:2 with up to 10 conservative amino acid substitutions.

has the following physicochemical properties (1) and (2):

(1) Action

i) The enzyme produces ketone by oxidizing alcohol using oxidized form of βnicotinamide adenine dinucleotide as a coenzyme, and

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The enzyme produces alcohol by reducing ketone using reduced form of βnicotinamide adenine dinucleotide as a coenzyme, and

## (2) Substrate specificity

- The enzyme preferentially oxidizes (R) 2 octanol of two optical isomers of 2 octanol, and
- The enzyme produces (S) 4-halo-3-hydroxybutyric acid esters by reducing 4haloacetoacetic acid esters.
- 16. (Currently amended) The method of claim 15, wherein the microorganism is a transformant comprising a recombinant vector into which a polynucleotide encoding the (R)-2octanol dehydrogenase is inserted.
  - 17. (Canceled)
- 18. (Currently amended) The method of claim 15 17, wherein the 4-haloacetoacetic acid ester derivative is 4-chloroacetoacetic acid ethyl ester and wherein the alcohol-(S)-4-halo-3hydroxybutyric acid ester derivative is (S)-4-chloro-3-hydroxybutyric acid ethyl ester.
  - 19. (Canceled)
  - 20. (Canceled)
- 21. (Currently amended) The method of claim 15, the method further comprising converting an oxidized form of  $\beta$ -nicotinamide adenine dinucleotide into a reduced form thereof.
  - 22. (Canceled)
  - 23. (Canceled)
  - 24. (Canceled)

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Please add new claims 25-38:

25. (New) The method of claim 15, wherein the (R)-2-octanol dehydrogenase has an optimal pH for the reduction reaction in a range from 5.0 to 6.5.

- 26. (New) The method of claim 15, wherein the reacting step is carried out with the microorganism producing the (R)-2-octanol dehydrogenase, and said microorganism belongs to the genus Candida or the genus Ogataea.
- 27. (New) The method of claim 15, wherein the reacting step is carried out with the microorganism producing the (R)-2-octanol dehydrogenase, and the microorganism belongs to the genus Pichia.
- 28. (New) The method of claim 15, wherein the (R)-2-octanol dehydrogenase is substantially pure, chemically treated, or in a cell-free extract.
- 29. (New) The method of claim 15, further comprising using a reduced form of βnicotinamide adenine dinucleotide (NADH) as a coenzyme.
- 30. (New) The method of claim 15, wherein the (R)-2-octanol dehydrogenase is encoded by a polynucleotide that hybridizes under stringent conditions with a nucleic acid probe consisting of the complement of the nucleotide sequence of SEQ ID NO: 1.
- 31. (New) The method of claim 15, wherein the (R)-2-octanol dehydrogenase comprises an amino acid sequence at least 70% identical to the amino acid sequence of SEO ID NO: 2.

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32. (New) The method of claim 31, wherein the (R)-2-octanol dehydrogenase comprises an amino acid sequence at least 80% identical to the amino acid sequence of SEQ ID NO: 2.

- 33. (New) The method of claim 31, wherein the (R)-2-octanol dehydrogenase comprises an amino acid sequence at least 90% identical to the amino acid sequence of SEQ ID NO: 2.
- 34. (New) The method of claim 31, wherein the (R)-2-octanol dehydrogenase comprises an amino acid sequence at least 95% identical to the amino acid sequence of SEQ ID NO: 2.
- 35. (New) The method of claim 15, wherein the (R)-2-octanol dehydrogenase comprises the amino acid sequence of SEQ ID NO: 2.
- 36. (New) The method of claim 35, wherein the (R)-2-octanol dehydrogenase consists of the amino acid sequence of SEQ ID NO: 2.
- 37. (New) The method of claim 15, wherein the (R)-2-octanol dehydrogenase comprises the amino acid sequence of SEQ ID NO:2 with up to 50 conservative amino acid substitutions.
- 38. (New) The method of claim 15, wherein the (R)-2-octanol dehydrogenase comprises the amino acid sequence of SEQ ID NO:2 with up to 10 conservative amino acid substitutions.